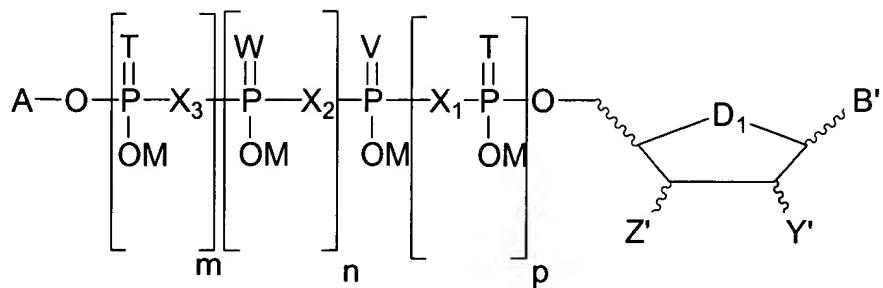


**THE AMENDMENTS**

**In the Claims**

1. (Previously Presented) A pharmaceutical formulation comprising a compound of general formula I, or salts thereof:

Formula I



wherein:

X<sub>1</sub>, X<sub>2</sub>, and X<sub>3</sub> are independently selected from the group consisting of oxygen, methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido; T, W, and V are independently oxygen or sulfur;

m= 0, 1 or 2;

n= 0, 1, or 2;

p= 0, 1, or 2 ;

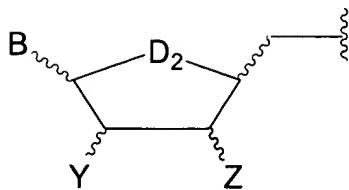
M= H or a pharmaceutically-acceptable inorganic or organic counterion;

D<sub>1</sub> =O or C;

B' is a purine or a pyrimidine residue according to general formulas IV and V which is linked to the 1' position of the furanose or carbocycle via the 9- or 1- position, respectively;

A is M or alkyl; or

A is a nucleoside residue which is defined as:



and which is linked to the phosphate chain via the 5' position of the furanose or carbocycle;  
wherein:

$D_2 = O$  or  $C$ ;

$B$  is a purine or a pyrimidine residue according to general formulas IV and V which is linked to the sugar or carbocycle via the 9- or 1- position, respectively;

wherein when  $D_1$  and  $D_2$  are oxygen, the furanose is in the  $\beta$ -configuration;

$Y' = H$ ,  $OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general Formula II or III;

$Z' = OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general Formula II or III;

$Z = OH$  or  $OR_3$ , where  $OR_3$  falls under the definition of general Formula II or III;

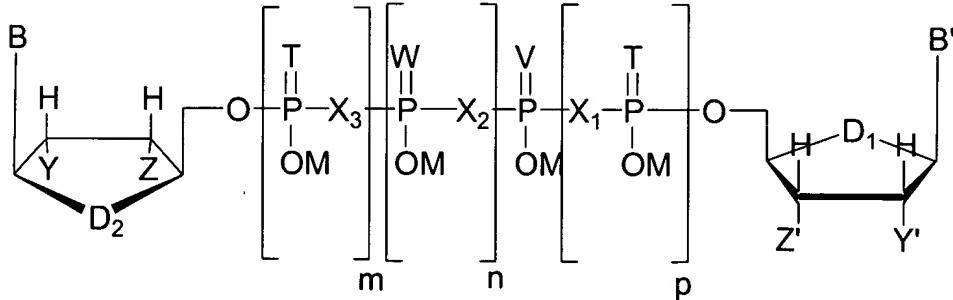
$Y = H$ ,  $OH$ , or  $OR_4$ , where  $OR_4$  falls under the definition of general Formula II or III;

provided that at least one of  $Y'$ ,  $Z'$ ,  $Z$ , and  $Y$  is  $OR_1$ ,  $OR_2$ ,  $OR_3$ , or  $OR_4$ , respectively;

wherein compounds of general Formula I are molecules whose structures fall within the definitions of Formula Ia and Formula Ib:

Formula Ia

wherein:



$X_1$ ,  $X_2$ , and  $X_3 = O$ ;

$T$ ,  $V$ , and  $W = O$ ;

$M = H$ ,  $NH_4^+$ ,  $Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion;

$Y' = H$ ,  $OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general formula II;

$Z' = OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general formula II;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula II;  
Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula II;  
provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;  
D<sub>1</sub> =O;  
D<sub>2</sub> =O or C;  
B and B' are purine or pyrimidine residues according to general formulas IV and V;  
m and p= 0, 1 or 2;  
n= 0 or 1;  
such that the sum of m+n+p is from 0 to 5; or

X<sub>1</sub>, X<sub>2</sub>, and X<sub>3</sub>=O;  
T, V, and W= O;  
M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion ;  
Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;  
Z'= OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;  
Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula III;  
Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula III;  
provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;  
D<sub>1</sub> =O;  
D<sub>2</sub> =O or C;  
B and B' are purine or pyrimidine residues according to general formulas IV and V;  
m and p= 0, 1 or 2;  
n= 0 or 1;  
such that the sum of m+n+p is from 0 to 5; or

X<sub>1</sub> and X<sub>3</sub>=O;  
X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene,  
dichloromethylene, monofluoromethylene, difluoromethylene, and imido;  
T, V, and W= O;  
M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion ;  
Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z' = OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;  
Z = OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula II;  
Y = H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula II;  
provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;  
D<sub>1</sub> = O;  
D<sub>2</sub> = O or C;  
B and B' are purine or pyrimidine residues according to general formulas IV and V;  
m and p = 0, 1 or 2;  
n = 1;  
such that the sum of m+n+p is from 0 to 5; or

X<sub>1</sub> and X<sub>3</sub> = O;  
X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene,  
dichloromethylene, monofluoromethylene, difluoromethylene, and imido;  
T, V, and W = O;  
M = H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion;  
Y' = H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;  
Z' = OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;  
Z = OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula III;  
Y = H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula III;  
provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

D<sub>1</sub> = O;  
D<sub>2</sub> is O or C;  
B and B' are purine or pyrimidine residues according to general formulas IV and V;  
m and p = 0, 1 or 2;  
n = 1;  
such that the sum of m+n+p is from 0 to 5; or

X<sub>1</sub> and X<sub>3</sub> = O;  
X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene,  
dichloromethylene, monofluoromethylene, difluoromethylene, and imido;  
T = S;

V and W=O;

M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula II;

Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula II;

provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

D<sub>1</sub> =O;

D<sub>2</sub> =O or C;

B and B' are purine or pyrimidine residues according to general formulas IV and V;

m, n, and p= 1; or

X<sub>1</sub> and X<sub>3</sub>=O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene,

dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T= S;

V and W=O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z'= OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula III;

Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula III;

provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

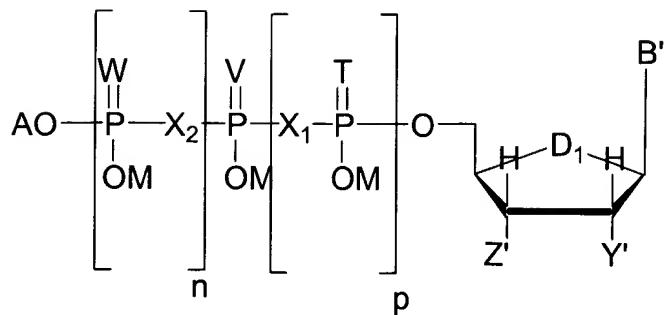
D<sub>1</sub> =O;

D<sub>2</sub> =O or C;

B and B' are purine or pyrimidine residues according to general formulas IV and V;

m, n, and p= 1;

Formula Ib



wherein:

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T, V, and W= O;

M= H, NH<sub>4</sub><sup>+</sup>, Na or other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

with the provision that at least one of Y' and Z' is OR<sub>1</sub> or OR<sub>2</sub>;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

n and p are 0, 1, or 2 such that the sum of n+p is from 1 to 3; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T, V, and W= O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z'= OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

n and p are 0, 1, or 2 such that the sum of n+p is from 1 to 3; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T and V= O;

W=S;

M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

with the provision that at least one of Y' and Z' is OR<sub>1</sub> or OR<sub>2</sub>;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T and V= O;

W=S;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z'= OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub> =O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T, V, and W= O;

M is selected from the group consisting of H, NH<sub>4</sub>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

with the provision that at least one of Y' and Z' is OR<sub>1</sub> or OR<sub>2</sub>;

D<sub>1</sub> = O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub>=O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T, V, and W= O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z'= H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> = O or C;

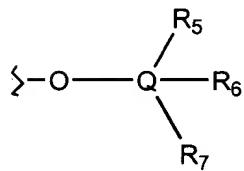
B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1;

wherein, for compounds according to Formula Ia or Ib, where Y'= OR<sub>1</sub>, Z'= OR<sub>2</sub>, Z= OR<sub>3</sub> and/or Y= OR<sub>4</sub>, at least one of the four is a residue which is linked directly to the corresponding 2' or 3' hydroxyl oxygen of the furanose or carbocycle via a carbon atom; wherein said residue falls within the scope of formula II or formula III:

Formula II



wherein:

O is the corresponding 2' or 3' oxygen of the furanose or carbocycle;

R<sub>5</sub>, R<sub>6</sub>, and R<sub>7</sub> are selected from the group consisting of H, an alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl, such that the moiety defined according to formula II is an ether; or

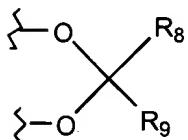
R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is selected from the group consisting of alkyl, cycloalkyl, aralkyl, and substituted aralkyl, such that the moiety defined according to formula II is an ester or thioester; or

R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is amino or mono- or disubstituted amino, where the substituents are selected from the group consisting of alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl, such that the moiety according to formula II is a carbamate or thiocarbamate; or

R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is selected from the group consisting of alkoxy, cycloalkoxy, aralkyloxy, aryloxy, substituted aralkyloxy, and substituted aryloxy, such that the moiety according to formula II is a carbonate or thiocarbonate; or

R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q and both the 2' and 3' oxygens of the furanose are directly bound to Q to form a cyclical carbonate or thiocarbonate, R<sub>7</sub> is not present;

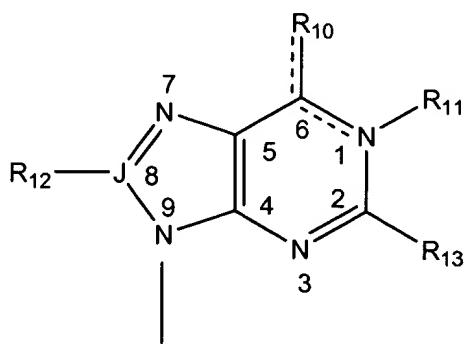
Formula III



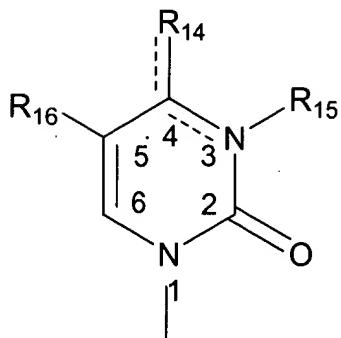
wherein:

O is the 2' and 3' oxygens of the furanose or carbocycle; and  
the 2' and 3' oxygens of the furanose or carbocycle are linked by a common carbon atom to form  
a cyclical acetal, cyclical ketal, or cyclical orthoester; and  
for cyclical acetals and ketals, R<sub>8</sub> and R<sub>9</sub> are independently selected from the group consisting of  
hydrogen, alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl; or are joined  
together to form a homocyclic or heterocyclic ring composed of 3 to 8 atoms, or  
for cyclical orthoesters, R<sub>8</sub> is selected from the group consisting of hydrogen, alkyl, cycloalkyl,  
aralkyl, aryl, substituted aralkyl, and substituted aryl,  
and R<sub>9</sub> is selected from the group consisting of alkyloxy, cycloalkyloxy, aralkyloxy, aryloxy,  
substituted aralkyloxy, and substituted aryloxy;  
B and B' are independently a purine residue, as in formula IV, linked through the 9- position, or a  
pyrimidine residue, as in formula V, linked through the 1- position;  
wherein, provided when D<sub>1</sub> and D<sub>2</sub> are oxygen, the ribosyl moieties are in the D- configuration;

Formula IV



Formula V



wherein:

R<sub>10</sub> and R<sub>14</sub> are selected from the group consisting of hydroxy, oxo, amino, mercapto, alkylthio, alkyloxy, aryloxy, alkylamino, cycloalkylamino, aralkylamino, arylamino, diaralkylamino, diarylamino, and dialkylamino, where the alkyl groups are optionally linked to form a heterocycle; or

R<sub>10</sub> and R<sub>14</sub> are acylamino according to Formula VI, provided that they incorporate an amino residue from the C-6 position of the purine or the C-4 position of the pyrimidine; or when R<sub>10</sub> in a purine or R<sub>14</sub> in a pyrimidine has as its first atom nitrogen, R<sub>10</sub> and R<sub>11</sub> or R<sub>14</sub> and R<sub>15</sub> are taken together to form a 5-membered fused imidazole ring, optionally substituted on the etheno ring with R<sub>5</sub>-R<sub>9</sub> selected from the group consisting of alkyl, cycloalkyl, aralkyl, or aryl moieties, as described above;

J is carbon or nitrogen, with the provision that when nitrogen, R<sub>12</sub> is not present;

R<sub>11</sub> is hydrogen, O or is absent;

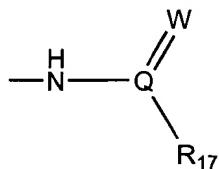
R<sub>12</sub> is selected from the group consisting of hydrogen, alkyl, azido, alkylamino, arylamino, aralkylamino, alkoxy, aryloxy, aralkyloxy, alkylthio, arythio, aralkylthio, and  $\omega$ -A(C<sub>1-6</sub>alkyl)B- wherein A and B are selected from the group consisting of independently amino, mercapto, hydroxy and carboxyl;

R<sub>13</sub> is selected from the group consisting of hydrogen, chlorine, amino, monosubstituted amino, disubstituted amino, alkylthio, arylthio, and aralkylthio, where the substituent on sulfur contains up to a maximum of 20 carbon atoms, with or without unsaturation;

R<sub>15</sub> is selected from the group consisting of hydrogen, and acyl, such as acetyl, benzoyl, phenylacetyl, with or without substituents;

$R_{16}$  is selected from the group consisting of hydrogen, methyl, alkyl, halo, alkyl, alkenyl, substituted alkenyl, alkynyl, and substituted alkynyl;

Formula VI



wherein:

NH is the amino residue at the C-6 position in a purine or the amino residue at the C-4 position in a pyrimidine;

Q is a carbon atom;

W is oxygen or sulfur;

$R_{17}$  is amino or mono- or disubstituted amino such that the moiety according to formula VI is a urea or thiourea; or

$R_{17}$  is selected from the group consisting of alkoxy, aralkyloxy, aryloxy, substituted aralkyloxy, and substituted aryloxy, such that the moiety according to formula VI is a carbamate or thiocarbamate; or

$R_{17}$  is selected from the group consisting of alkyl, cycloalkyl, aralkyl, and aryl, with or without substituents or heteroatoms, such that the moiety according to formula VI is an amide.

2. (Currently Amended) The compound according to Claim 37 38, wherein said compound is fluorescently labeled and used as a biochemical probe for the P2<sub>T</sub> receptor.

3. (Previously Presented) A method of preventing or treating diseases or conditions associated with platelet aggregation comprising:

administering to a patient a pharmaceutical formulation according to Claim 1, wherein said compound is effective to bind the P2<sub>T</sub>

receptors on platelets and inhibit ADP-induced platelet aggregation.

4. (Canceled).

5. (Previously Presented) The method according to Claim 3, wherein said pharmaceutical formulation is administered to reduce the incidence of dose-related adverse side effects of other therapeutic agents used to prevent, manage or treat platelet aggregation disorders.

6. (Previously Presented) The method according to Claim 3, wherein said administering is systemic administration of said compound.

7. (Original) The method according to Claim 6, wherein said systemic administration is administration of an injectable form of said compound, such that a therapeutically effective amount of said compound contacts the target platelets of said patient via systemic absorption and circulation.

8. (Previously Presented) The method according to Claim 6, wherein said systemic administration is accomplished by administering an oral form of said compound, such that a therapeutically effective amount of said compound contacts the target platelets of said patient via systemic absorption and circulation.

9. (Original) The method according to Claim 6, wherein said systemic administration is administration of said compound in a form of a transdermal patch or a transdermal pad, such that a therapeutically effective amount of said compound contacts the target platelets of said patient via systemic absorption and circulation.

10. (Original) The method according to Claim 6, wherein said systemic administration is administration of a liquid/liquid suspension of said compound via nose drops or nasal spray, or administration of a nebulized liquid to oral or nasopharyngeal airways of said subject, such that a therapeutically effective amount of said compound inhibits platelet aggregation.

11. (Original) The method according to Claim 6, wherein said systemic administration comprises infusion of said compound to target platelets via a device selected from a group consisting of a pump catheter system and a continuous or selective release device.
12. (Original) The method according to Claim 6, wherein said systemic administration is administration of a suppository form of said compound, such that a therapeutically effective amount of said compound contacts the target platelets of said patient via systemic absorption and circulation.
13. (Original) The method according to Claim 6, wherein said systemic administration is vaginal administration in dosage unit formulations containing conventional non-toxic pharmaceutically acceptable carriers, adjuvants and vehicles.
14. (Original) The method according to Claim 6, wherein said compound is administered to a patient by an intravitreal delivery.
15. (Original) The method according to Claim 6, wherein said systemic administration is administration of an intra-operative instillation of a gel, cream, powder, foam, crystals, liposomes, spray or liquid suspension form of said compound, such that a therapeutically effective amount of said compound contracts the target platelets of said patient via systemic absorption and circulation.
16. (Previously Presented) The method according to Claim 3, wherein said diseases or conditions associated with platelet aggregation are disorders or procedures characterized by thrombosis, primary arterial thrombotic complications of atherosclerotic disease, thrombotic complications of interventions of atherosclerotic disease, thrombotic complications of surgical or mechanical damage, mechanically – induced platelet activation, shunt occlusion, thrombosis secondary to vascular damage and inflammation, indications with a diffuse thrombotic/platelet consumption component, venous thrombosis, coronary arterial thrombosis, pathological effects of atherosclerosis and arteriosclerosis, platelet aggregation and clot formation in blood and blood products during storage, chronic or acute states of hyper-aggregability, reocclusion of an artery or vein following fibrinolytic therapy, platelet adhesion associated with extracorporeal circulation,

thrombotic complications associated with thrombolytic therapy, thrombotic complications associated with coronary and other angioplasty, or thrombotic complications associated with coronary artery bypass procedures.

17. (Previously Presented) The method according to Claim 16, wherein said disorders or procedures characterized with thrombosis are unstable angina, coronary angioplasty, or myocardial infarction.

18. (Previously Presented) The method according to Claim 16, wherein said primary arterial thrombotic complications of atherosclerosis are thrombotic stroke, peripheral vascular disease, or myocardial infarction without thrombolysis.

19. (Previously Presented) The method according to Claim 16, wherein said thrombotic complications of interventions of atherosclerotic disease are associated with angioplasty, endarterectomy, stent placement, coronary or other vascular graft surgery.

20. (Previously Presented) The method according to Claim 16, wherein said thrombotic complications of surgical or mechanical damage are associated with tissue salvage following surgical or accidental trauma, reconstructive surgery including skin flaps, or reductive surgery such as breast reduction.

21. (Previously Presented) The method according to Claim 16, wherein said mechanically – induced platelet activation is caused by cardiopulmonary bypass resulting in microthromboembolism.

22. (Previously Presented) The method according to Claim 16, wherein said shunt occlusion is renal dialysis or plasmapheresis.

23. (Previously Presented) The method according to Claim 16, wherein said thrombosis secondary to vascular damage and inflammation is vasculitis, arteritis, glomerulonephritis or organ graft rejection.
24. (Previously Presented) The method according to Claim 16, wherein said indications with a diffuse thrombotic/platelet consumption component are disseminated intravascular coagulation, thrombotic thrombocytopenic purpura, hemolytic uremic syndrome, heparin-induced thrombocytopenia, or pre-eclampsia/eclampsia.
25. (Previously Presented) The method according to Claim 16, wherein said venous thrombosis is deep vein thrombosis, veno-occlusive disease, hematological conditions, or migraine.
26. (Previously Presented) The method according to Claim 25, wherein said hematological conditions are thrombocythemia or polycythemia.
27. (Previously Presented) The method according to Claim 16, wherein said coronary arterial thrombosis is associated with unstable angina, coronary angioplasty or acute myocardial infarction.
28. (Previously Presented) The method according to Claim 16, wherein pathological effects of atherosclerosis and arteriosclerosis are arteriosclerosis, acute myocardial infarction, chronic stable angina, unstable angina, transient ischemic attacks, strokes, peripheral vascular disease, arterial thrombosis, preeclampsia, embolism, restenosis or abrupt closure following angioplasty, carotid endarterectomy, or anastomosis of vascular grafts.

29. (Previously Presented) The method according to Claim 16, wherein said chronic or acute states of hyper-aggregability is caused by DIC, septicemia, surgical or infectious shock, post-operative and post-partum trauma, cardiopulmonary bypass surgery, incompatible blood transfusion, abruptio placenta, thrombotic thrombocytopenic purpura, snake venom or immune diseases.
30. (Original) The method according to Claim 16, wherein said reocclusion of an artery or vein following fibrinolytic therapy is inhibited by internal administration of said compound with a fibrinolytic agent.
31. (Previously Presented) The method according to Claim 30, wherein said fibrinolytic agent is selected from the group consisting of natural or synthetic products which directly or indirectly cause lysis of a fibrin clot.
32. (Previously Presented) The method according to Claim 30, wherein said fibrinolytic agent is a plasminogen activator selected from the group consisting of anistreplase, urokinase, pro-urokinase, streptokinase, tissue plasminogen activator and mutants or variants thereof, which retain plasminogen activator activity.
33. (Previously Presented) The method according to Claim 32, wherein said variants are selected from the group consisting of variants which have been chemically modified, variants which one or more amino acids have been added, deleted or substituted and variants with one or more modified functional domains.

34. (Previously Presented) The method according to Claim 33, wherein said modified functional domains are added, deleted or altered by combining the active site of one plasminogen activator or fibrin binding domain with another plasminogen activator or fibrin binding molecule.

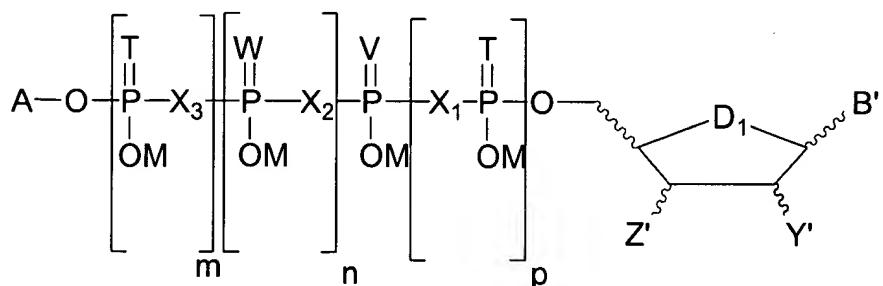
35. (Previously Presented) The pharmaceutical formulation according to Claim 1, wherein said formulation is sterile.

36. (Previously Presented) The pharmaceutical formulation according to Claim 1, wherein said formulation further comprises a pharmaceutical carrier.

37. (Previously Presented) The pharmaceutical formulation according to Claim 1, wherein said formulation further comprises a buffering agent.

38. (Previously Presented) A compound of general formula I, or salts thereof:

Formula I



wherein:

X<sub>1</sub>, X<sub>2</sub>, and X<sub>3</sub> are independently selected from the group consisting of oxygen, methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido; T, W, and V are independently oxygen or sulfur;

m= 0, 1 or 2;

n= 0, 1, or 2;

p= 0, 1, or 2 ;

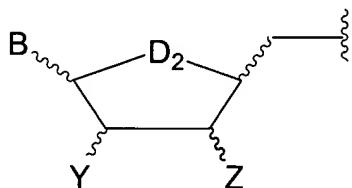
M= H or a pharmaceutically-acceptable inorganic or organic counterion;

D<sub>1</sub> =O or C;

B' is a purine or a pyrimidine residue according to general formulas IV and V which is linked to the 1' position of the furanose or carbocycle via the 9- or 1- position, respectively;

A is M or alkyl; or

A is a nucleoside residue which is defined as:



and which is linked to the phosphate chain via the 5' position of the furanose or carbocycle;  
wherein:

D<sub>2</sub> =O or C;

B is a purine or a pyrimidine residue according to general formulas IV and V which is linked to the sugar or carbocycle via the 9- or 1- position, respectively;

wherein when D<sub>1</sub> and D<sub>2</sub> are oxygen, the furanose is in the  $\beta$ -configuration;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general Formula II or III;

Z'= OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general Formula II or III;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general Formula II or III;

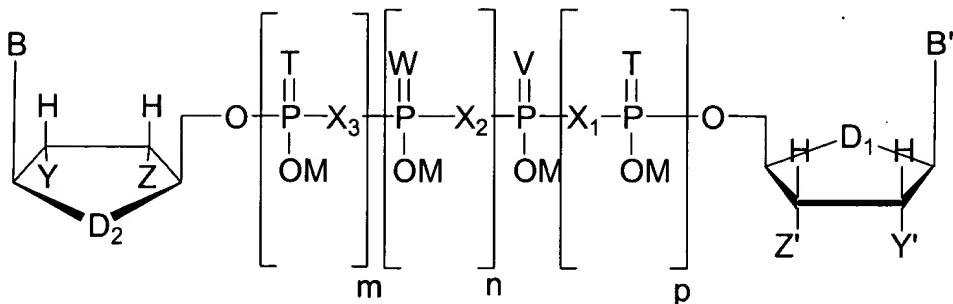
Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general Formula II or III;

provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

wherein compounds of general Formula I are molecules whose structures fall within the definitions of Formula Ia and Formula Ib:

Formula Ia

wherein:



$X_1, X_2$ , and  $X_3=O$ ;

$T, V$ , and  $W=O$ ;

$M=H, NH_4^+, Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion;

$Y'=H, OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general formula II;

$Z'=OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general formula II;

$Z=OH$  or  $OR_3$ , where  $OR_3$  falls under the definition of general formula II;

$Y=H, OH$ , or  $OR_4$ , where  $OR_4$  falls under the definition of general formula II;

provided that at least one of  $Y', Z', Z$ , and  $Y$  is  $OR_1, OR_2, OR_3$ , or  $OR_4$ , respectively;

$D_1=O$ ;

$D_2=O$  or  $C$ ;

$B$  and  $B'$  are purine or pyrimidine residues according to general formulas IV and V;

$m$  and  $p=0, 1$  or  $2$ ;

$n=0$  or  $1$ ;

such that the sum of  $m+n+p$  is from  $0$  to  $5$ ; or

$X_1, X_2$ , and  $X_3=O$ ;

$T, V$ , and  $W=O$ ;

$M=H, NH_4^+, Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion ;

$D_1=O$ ;  $Y'=H, OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general formula III;

$Z'=OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general formula III;

$Z=OH$  or  $OR_3$ , where  $OR_3$  falls under the definition of general formula III;

$Y=H, OH$ , or  $OR_4$ , where  $OR_4$  falls under the definition of general formula III;

provided that at least one of  $Y', Z', Z$ , and  $Y$  is  $OR_1, OR_2, OR_3$ , or  $OR_4$ , respectively;

$D_2 = O$  or  $C$ ;

$B$  and  $B'$  are purine or pyrimidine residues according to general formulas IV and V;

$m$  and  $p = 0, 1$  or  $2$ ;

$n = 0$  or  $1$ ;

such that the sum of  $m+n+p$  is from  $0$  to  $5$ ; or

$X_1$  and  $X_3 = O$ ;

$X_2$  is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

$T, V$ , and  $W = O$ ;

$M = H, NH_4^+, Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion;

$Y' = H, OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general formula II;

$Z' = OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general formula II;

$Z = OH$  or  $OR_3$ , where  $OR_3$  falls under the definition of general formula II;

$Y = H, OH$ , or  $OR_4$ , where  $OR_4$  falls under the definition of general formula II;

provided that at least one of  $Y', Z', Z$ , and  $Y$  is  $OR_1, OR_2, OR_3$ , or  $OR_4$ , respectively;

$D_1 = O$ ;

$D_2 = O$  or  $C$ ;

$B$  and  $B'$  are purine or pyrimidine residues according to general formulas IV and V;

$m$  and  $p = 0, 1$  or  $2$ ;

$n = 1$ ;

such that the sum of  $m+n+p$  is from  $0$  to  $5$ ; or

$X_1$  and  $X_3 = O$ ;

$X_2$  is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

$T, V$ , and  $W = O$ ;

$M = H, NH_4^+, Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion;

$Y' = H, OH$ , or  $OR_1$ , where  $OR_1$  falls under the definition of general formula III;

$Z' = OH$  or  $OR_2$ , where  $OR_2$  falls under the definition of general formula III;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula III;  
Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula III;

provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

D<sub>1</sub> =O;

D<sub>2</sub> is O or C;

B and B' are purine or pyrimidine residues according to general formulas IV and V;

m and p= 0,1 or 2;

n=1;

such that the sum of m+n+p is from 0 to 5; or

X<sub>1</sub> and X<sub>3</sub>=O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T= S;

V and W=O;

M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

Z= OH or OR<sub>3</sub>, where OR<sub>3</sub> falls under the definition of general formula II;

Y= H, OH, or OR<sub>4</sub>, where OR<sub>4</sub> falls under the definition of general formula II;

provided that at least one of Y', Z', Z, and Y is OR<sub>1</sub>, OR<sub>2</sub>, OR<sub>3</sub>, or OR<sub>4</sub>, respectively;

D<sub>1</sub> =O;

D<sub>2</sub> =O or C;

B and B' are purine or pyrimidine residues according to general formulas IV and V;

m, n, and p= 1; or

X<sub>1</sub> and X<sub>3</sub>=O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T= S;

V and W=O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable

inorganic or organic counterion;

$Y' = H, OH, \text{ or } OR_1$ , where  $OR_1$  falls under the definition of general formula III;

$Z' = OH \text{ or } OR_2$ , where  $OR_2$  falls under the definition of general formula III;

$Z = OH \text{ or } OR_3$ , where  $OR_3$  falls under the definition of general formula III;

$Y = H, OH, \text{ or } OR_4$ , where  $OR_4$  falls under the definition of general formula III;

provided that at least one of  $Y'$ ,  $Z'$ ,  $Z$ , and  $Y$  is  $OR_1$ ,  $OR_2$ ,  $OR_3$ , or  $OR_4$ , respectively;

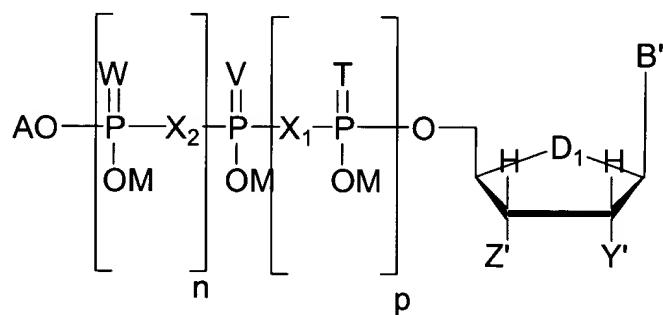
$D_1 = O$ ;

$D_2 = O \text{ or } C$ ;

$B$  and  $B'$  are purine or pyrimidine residues according to general formulas IV and V;

$m, n, \text{ and } p = 1$ ;

Formula Ib



wherein:

$A$  is  $M$  or alkyl;

$X_1$  and  $X_2 = O$ ;

$T, V, \text{ and } W = O$ ;

$M = H, NH_4^+, Na^+$  or other pharmaceutically-acceptable inorganic or organic counterion;

$Y' = H, OH, \text{ or } OR_1$ , where  $OR_1$  falls under the definition of general formula II;

$Z' = H, OH \text{ or } OR_2$ , where  $OR_2$  falls under the definition of general formula II;

with the provision that at least one of  $Y'$  and  $Z'$  is  $OR_1$  or  $OR_2$ ;

$D_1 = O \text{ or } C$ ;

$B'$  is purine or pyrimidine residue according to general formulas IV and V;

$n$  and  $p$  are 0, 1, or 2 such that the sum of  $n+p$  is from 1 to 3; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T, V, and W= O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= OR<sub>1</sub>, where R<sub>1</sub> falls under the definition of general formula III;

Z'= OR<sub>2</sub>, where R<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

n and p are 0, 1, or 2 such that the sum of n+p is from 1 to 3; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T and V= O;

W=S;

M= H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> or other pharmaceutically-acceptable inorganic or organic counterion;

Y'= H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z'= H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

with the provision that at least one of Y' and Z' is OR<sub>1</sub> or OR<sub>2</sub>;

D<sub>1</sub> =O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub> and X<sub>2</sub>= O;

T and V= O;

W=S;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y'= OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z' = OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> = O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub> = O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T, V, and W = O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

Y' = H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula II;

Z' = H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula II;

with the provision that at least one of Y' and Z' is OR<sub>1</sub> or OR<sub>2</sub>;

D<sub>1</sub> = O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;

p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;

n=1; or

A is M or alkyl;

X<sub>1</sub> = O;

X<sub>2</sub> is selected from the group consisting of methylene, monochloromethylene, dichloromethylene, monofluoromethylene, difluoromethylene, and imido;

T, V, and W = O;

M is selected from the group consisting of H, NH<sub>4</sub><sup>+</sup>, Na<sup>+</sup> and other pharmaceutically-acceptable inorganic or organic counterion;

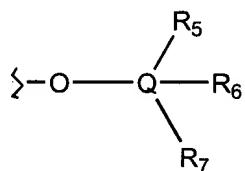
Y' = H, OH, or OR<sub>1</sub>, where OR<sub>1</sub> falls under the definition of general formula III;

Z' = H, OH or OR<sub>2</sub>, where OR<sub>2</sub> falls under the definition of general formula III;

D<sub>1</sub> = O or C;

B' is purine or pyrimidine residue according to general formulas IV and V;  
p is 0, 1, or 2 such that the sum of n+p is from 1 to 3;  
n=1;  
wherein, for compounds according to Formula Ia or Ib, where Y'=OR<sub>1</sub>, Z'=OR<sub>2</sub>, Z=OR<sub>3</sub> and/or Y=OR<sub>4</sub>, at least one of the four is a residue which is linked directly to the corresponding 2' or 3' hydroxyl oxygen of the furanose or carbocycle via a carbon atom; wherein said residue falls within the scope of formula II or formula III:

Formula II



wherein:

O is the corresponding 2' or 3' oxygen of the furanose or carbocycle;

R<sub>5</sub>, R<sub>6</sub>, and R<sub>7</sub> are selected from the group consisting of H, an alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl, such that the moiety defined according to formula II is an ether; or

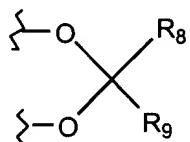
R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is selected from the group consisting of alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl, such that the moiety defined according to formula II is an ester or thioester; or

R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is amino or mono- or disubstituted amino, where the substituents are selected from the group consisting of alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl, such that the moiety according to formula II is a carbamate or thiocarbamate; or

R<sub>5</sub> and R<sub>6</sub> are taken together to be oxygen or sulfur doubly bonded to Q, and R<sub>7</sub> is selected from the group consisting of alkoxy, cycloalkoxy, aralkyloxy, aryloxy, substituted aralkyloxy, and substituted aryloxy, such that the moiety according to formula II is a carbonate or thiocarbonate; or

$R_5$  and  $R_6$  are taken together to be oxygen or sulfur doubly bonded to Q and both the 2' and 3' oxygens of the furanose are directly bound to Q to form a cyclical carbonate or thiocarbonate,  $R_7$  is not present;

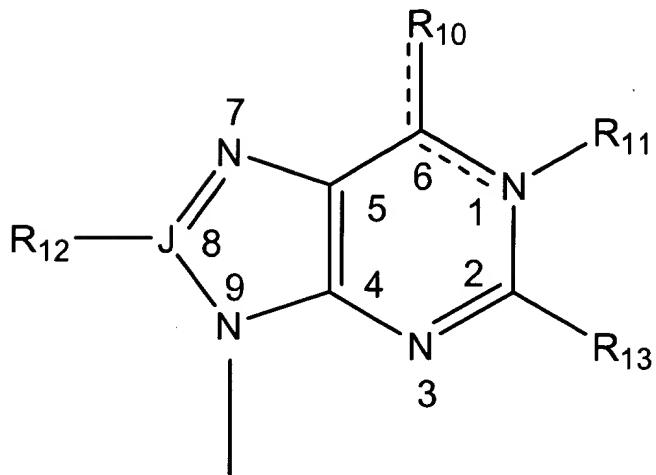
Formula III



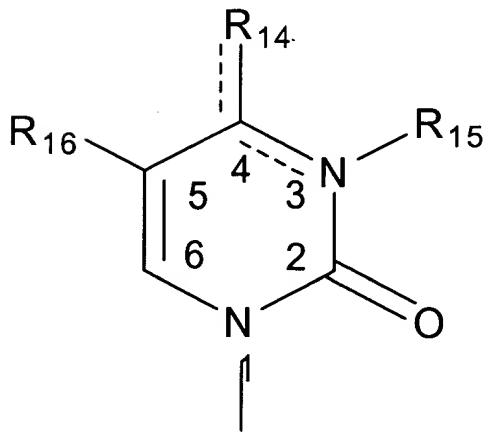
wherein:

O is the 2' and 3' oxygens of the furanose or carbocycle; and  
the 2' and 3' oxygens of the furanose or carbocycle are linked by a common carbon atom to form a cyclical acetal, cyclical ketal, or cyclical orthoester; and  
for cyclical acetals and ketals,  $R_8$  and  $R_9$  are independently selected from the group consisting of hydrogen, alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl; or are joined together to form a homocyclic or heterocyclic ring composed of 3 to 8 atoms, or  
for cyclical orthoesters,  $R_8$  is selected from the group consisting of hydrogen, alkyl, cycloalkyl, aralkyl, aryl, substituted aralkyl, and substituted aryl,  
and  $R_9$  is selected from the group consisting of alkyloxy, cycloalkyloxy, aralkyloxy, aryloxy, substituted aralkyloxy, and substituted aryloxy;  
B and B' are independently a purine residue, as in formula IV, linked through the 9- position, or a pyrimidine residue, as in formula V, linked through the 1- position;  
wherein, provided when  $D_1$  and  $D_2$  are oxygen, the ribosyl moieties are in the D- configuration;

Formula IV



Formula V



wherein:

$R_{10}$  and  $R_{14}$  are selected from the group consisting of alkylthio, alkyloxy, aryloxy, cycloalkylamino, aralkylamino, arylamino, diaralkylamino, and diarylamino, where the alkyl groups are optionally linked to form a heterocycle; or

$R_{10}$  and  $R_{14}$  are acylamino according to Formula VI, provided that they incorporate an amino residue from the C-6 position of the purine or the C-4 position of the pyrimidine; or

when  $R_{10}$  in a purine or  $R_{14}$  in a pyrimidine has as its first atom nitrogen,  $R_{10}$  and  $R_{11}$  or  $R_{14}$  and  $R_{15}$  are taken together to form a 5-membered fused imidazole ring, optionally substituted on the etheno ring with  $R_5$ - $R_9$  selected from the group consisting of alkyl, cycloalkyl, aralkyl, or aryl moieties, as described above;

$J$  is carbon or nitrogen, with the provision that when nitrogen,  $R_{12}$  is not present;

R<sub>11</sub> is hydrogen, O or is absent;

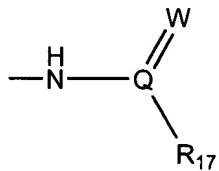
R<sub>12</sub> is selected from the group consisting of hydrogen, alkyl, azido, alkylamino, arylamino, aralkylamino, alkoxy, aryloxy, aralkyloxy, alkylthio, arythio, aralkylthio, and  $\omega$ -A(C<sub>1-6</sub>alkyl)B- wherein A and B are selected from the group consisting of independently amino, mercapto, hydroxy and carboxyl;

R<sub>13</sub> is selected from the group consisting of hydrogen, chlorine, amino, monosubstituted amino, disubstituted amino, alkylthio, arylthio, and aralkylthio, where the substituent on sulfur contains up to a maximum of 20 carbon atoms, with or without unsaturation;

R<sub>15</sub> is selected from the group consisting of hydrogen, and acyl, such as acetyl, benzoyl, phenylacetyl, with or without substituents;

R<sub>16</sub> is selected from the group consisting of hydrogen, methyl, alkyl, halo, alkyl, alkenyl, substituted alkenyl, alkynyl, and substituted alkynyl;

Formula VI



wherein:

NH is the amino residue at the C-6 position in a purine or the amino residue at the C-4 position in a pyrimidine;

Q is a carbon atom;

W is oxygen or sulfur;

R<sub>17</sub> is amino or mono- or disubstituted amino such that the moiety according to formula VI is a urea or thiourea; or

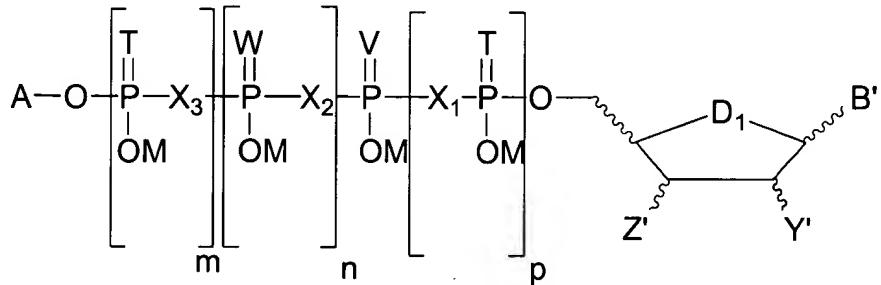
R<sub>17</sub> is selected from the group consisting of alkoxy, aralkyloxy, aryloxy, substituted aralkyloxy, and substituted aryloxy, such that the moiety according to formula VI is a carbamate or thiocarbamate; or

R<sub>17</sub> is selected from the group consisting of alkyl, cycloalkyl, aralkyl, and aryl, with or without

substituents or heteroatoms, such that the moiety according to formula VI is an amide.

39. (Previous Presented) The pharmaceutical formulation of Claim 1 wherein said compound is a compound of Formula I:

Formula I



wherein:

V = O;

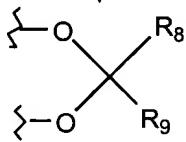
n = m = p = 0;

A = M;

M = H or a pharmaceutically-acceptable inorganic or organic counterion;

D<sub>1</sub> = O;

Formula III



wherein:

O is the 2' and 3' oxygens of the furanose; and

the 2' and 3' oxygens of the furanose are linked by a common carbon atom to form a cyclical acetal; and

R<sub>8</sub> is hydrogen; and

R<sub>9</sub> is selected from the group consisting of aralkyl, aryl, substituted aralkyl, and substituted aryl;

in which the aralkyl groups are from 1 to 5 carbons in the alkyl portion, and are: monocyclic moieties from 4 to 8 carbons without heteroatoms in the aryl portion; and the aryl groups are monocyclic moieties from 4 to 8 carbons, without heteroatoms;

B' is a purine residue according to general Formula IV

wherein:

R<sub>10</sub> is acylamino, according to Formula VI;

W is oxygen; and

R<sub>17</sub> is amino or mono- or disubstituted amino such that the moiety according to Formula VI is a urea;

J = carbon;

R<sub>11</sub> is absent;

R<sub>12</sub> is hydrogen; and

R<sub>13</sub> is hydrogen.